Docket No.: NOR-014CP4/286336.153US1

Reply dated February 25, 2008

## APPENDIX A

Attached is a Declaration Under 37 C.F.R. § 1.132 from Dr. Richard Boyd.

#### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application No.

10/749,122

Art Unit:

1632

Applicant:

Richard L. Boyd

Examiner:

David A. Montanari

Date Filed:

December 30, 2003

Conf. No.

3280

Docket No.

NOR-014CP4 and 286336.153US1

Cust. No.

23483

Title:

DISEASE PREVENTION BY REACTIVATION OF THE THYMUS

#### DECLARATION OF DR. RICHARD L. BOYD UNDER 37 C.F.R. § 1.132

Dear Sir:

In connection with the above-referenced patent application, I, Richard L. Boyd, declare as follows:

1. I received my Ph.D. degree in Immunology from Monash University, Australia in 1976. I was a Senior Tutor in the Department of Pathology and Immunology at Monash University from 1976-1977. I held a research fellowship at the University of Innsbruck, Austria, from 1978-1982, following which I held a Research Fellow position from 1983-1984. I was a Lecturer at Monash University from 1985-1994. Currently, I am an Associate Professor in the Department of Immunology at Monash University. I am also the Director of the Immunology Platform Program at the Australian Stem Cell Center and the Deputy Director of Monash Immunology and Stem Cell Laboratories. I have authored or co-authored more than 193 journal articles in the area of immunology and presented over 500 oral presentations at conferences and research institutes. I was the Editor-in-Chief of Developmental Immunology from 1999-2003, and a member of the editorial boards of Developmental and Comparative Immunology (1993-2003) and of Clinical and Developmental Immunology (2004-2006). I review articles for numerous journals including, Nature, Nature Immunology, Blood, Autoimmunity, and the Journal of Experimental Medicine. I was the recipient of two awards from the Australian Federal Government and a co-recipient of two international prizes for my research. My curriculum vitae, which includes a list of my publications and presentations, is provided as **Attachment A** following *page 6* of this Declaration.

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2. I am the Chief Scientific Officer of Norwood Immunology Limited, the Assignee of the

above-referenced application.

3. I am the sole inventor of the above-referenced patent application and accordingly am

familiar with the above-referenced patent application. I am also familiar with the Office Action

dated August 24, 2007 (hereinafter "Office Action") in the above-referenced application.

4. As I understand the Office Action, the Examiner rejected the pending claims, in part, as

not being enabling (see, Office Action, pages 4-7). The Office Action states, in relevant part, that:

"Whereas the nature of the invention is a method of increasing thymus activity (producing

more thymocytes) by disruption of sex steroid influences via chemical castration, the art teaches

that such a method would not prevent or treat all disease." (see, Office Action, page 4, last

paragraph).

5. As amended, the claims of my application are directed, in part, to methods for treating

or increasing resistance to a viral infection in a patient by reactivating the thymus of the patient.

My patent application teaches a trained clinical immunologist how to make and use the

invention as claimed in this application.

6. The application provides sufficient guidance to a trained clinical immunologist to

practice the invention as claimed, for example, at page 30, lines 4-30; page 60, lines 14-31;

Example 3; Figures 14-19; and Example 10 of the application as filed.

7. In this Declaration, I provide further experimental evidence to show that the disclosure

in the application as filed is sufficient to teach a trained clinical immunologist to make and use

my presently claimed invention. Specifically, the experiments described below show that the

claimed methods are useful in treating or increasing resistance to influenza infections.

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8. Cytotoxic T cells play a major role in the control and eventual elimination of viral infections. Subjects with decreased thymic activity (*e.g.*, the aged) display increased susceptibility to viral infections, such as influenza A virus, and this may reflect either diminished CTL capacity or loss of antigen-specific CTL. Utilizing a well-characterized model of influenza A virus infection in C57BL/6J mice, we assessed the capacity of aged mice to respond to both primary and secondary viral infection.

- 8. Intranasal infection of C57BL/6J (B6) mice with Influenza A virus causes an acute respiratory pneumonia with virus cleared around ten days post infection. This time point corresponds with the peak of the lymphocytic infiltrate found in broncheaveolar lavage (BAL). The primary CD8<sup>-</sup> T cell response is largely specific for three determinants derived from the nucleoprotein (NP366-374 (NP366); amino acid sequence ASNENMETM), acidic polymerase (PA224-233 (PA224); amino acid sequence SSLENFRAYV) and basic polymerase 1 (PB1703-711 (PB1703); amino acid sequence SSYRRYPVGI) of the virus. Influenza A virus-specific CTL play a major role in limiting and eventual clearance of influenza A virus infection. For example, increased CTL numbers after infection correlate with decreased viral shedding and protection from heterologous virus challenge where humoral immunity is limited.
- 9. The capacity of aged mice (9 mths 24 mths) to respond to influenza A virus infection, as measured by CTL responses against the epitopes NP366, PA224 and PB1703, was compared to young animals. Young mice (6-8 weeks) or mice aged 9, 18, or 24 mths were infected intranasally (i.n.) with  $10^4$  pfu of A/HKx31 influenza A virus. The specific CTL responses for NP366, PA224, and PB1703 were determined by intracellular cytokine staining at days 3, 7, 10, and 14 after infection. No difference was observed between the young and aged mice at day 3 after infection, presumably due to low CTL numbers that soon after infection. However, aged mice demonstrated both a lower percentage and number of CD8 $\alpha$ +IFN- $\gamma$ + CTL at day 7 and 10 (p<0.01) for all epitopes tested (*see*, **Attachment B**, Figs.1A, and 1B). These data show that

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following intransal influenza infection, aged mice have a decreased number of influenzaspecific CTLs compared with younger mice.

- 10. We next determined the naïve T cell:memory T cell ratio in the young and old mice. Using expression of CD62L and CD44 to define naïve (CD62Lhi/CD44lo) and memory (CD62Lhi/CD44lo) CTL phenotypes, we observed a significant age-dependent decline in the naïve T cell:memory CTL ratio (*see*, **Attachment C**, Fig 2A, p<0.01) when compared to young mice. This loss of naïve T cell precursors is presumably due to an aged-related decrease in thymic output. Sex steroid ablation (SSA) can increase thymic output due to release of thymic stroma from sex-steroid repression of thymic function. Utilizing castration as a method of SSA, the impact of removing sex-steroid repression on the naïve T cell:memory T cell ratio was determined in mice aged 9, 18, and 24 mths (*see*, **Attachment C**, Fig. 2B). A significant (p<0.05) increase in the naïve T cell:memory T cell ratio was observed in all aged mice compared to their sham castrated counterparts. However the level of restoration appeared age-dependent with 9 mth old castrated mice showing a return to a naive:memory ratio similar to that seen in young mice (*see*, **Attachment C**, Fig 2B). In contrast, despite a level of restoration, 18 mth and 24 mth old Cx mice did not show a return to the levels seen in young mice (p<0.01).
- 11. We then determined whether increased numbers of naive CD8 $\alpha^+$  T cells observed following castration would restore virus-specific CTL responses to influenza A virus infection. Mice aged 9, 18, and 24 mths were castrated and infected i.n. with with A/HKx31 6 weeks later. Fewer NP366-specific CTL were elicited after infection of sham castrated mice demonstrating the age related diminished response to infection compared to young mice (*see*, **Attachment D**, Fig. 3A, B, p<0.05). Following castration both 9 mth and 18 mth old mice showed recovery of both the proportion (*see*, **Attachment D**, Fig. 7A, p<0.05) and absolute number (Fig. 3B, p<0.05) of CD8 $\alpha^+$ IFN $\gamma^+$  CTL when compared that seen in young mice (*see*, **Attachment D**, Fig 3A, B). Interestingly, 24 mth old castrated mice did not show the same recovery indicating that sex

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steroid ablation may have age-limited positive effects on improving the response to influenza virus infection (*see*, **Attachment D**, Fig. 3A, B).

- 12. Analysis of CTL cytotoxicity demonstrated that the increase in CD8 $\alpha$ \*IFN $\gamma$ \* cell numbers due to castration corresponded with increased anti-influenza-specific cytotoxic activity in 9 mth old mice. Importantly, the restoration was equivalent to the cytotoxic responses observed in young mice while sham castrated mice demonstrated decreased cytoxicity compared to young mice (*see*, **Attachment D**, Fig. 3C). A strong correlation was again seen between cytotoxic activity and the number of NP366-specific CTL (*see*, **Attachment D**, Fig. 3D). Furthermore, there was no difference in the capacity of sham castrated and castrated aged mice to produce cytokine after peptide stimulation (data not shown).
- 13. To determine if the subsequent restoration of CTL responses improved viral clearance, lung viral titres were determined day 7 after i.n. infection of young mice and compared to aged sham castrated and castrated mice (*see*, **Attachment E**, Fig 4). Sham castrated mice aged 9 and 24 mths had significantly higher viral titres when compared young mice (p<0.05). Importantly, 9 mth castrated mice showed significantly improved viral lung clearance to a level similar to that observed in normal young mice (*see*, **Attachment E**, Fig 4, p<0.05). There was also improved viral clearance in castrated mice aged 24 mths although this wasn't to the level observed in 9 mth castrated mice. Overall these data show that sex-steroid ablation restores the numbers of anti-influenza A virus CTL generated after infection and that correlates in improved viral clearance.
- 14. In summary, these data show that reactivating the thymus of the patient can be used to treat or increase resistance to a viral infection.
- 15. Based on the discussion above, I respectfully submit that the above-referenced application does teach how to make and use the invention as presently claimed.

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16. I further declare that all statements made herein of my own knowledge are true and that

all statements made on information and belief are believed to be true and further that these

statements are made with the knowledge that willful false statements and the like so made are

punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States

Code and that such willful false statements may jeopardize the validity of the application or any

patent issuing thereon.

Date: February 22nd 2008

Richard L. Boyd, Ph.D.

Kichard Bayd.

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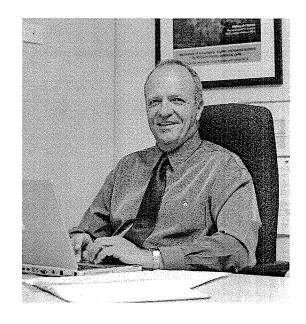
## ATTACHMENT A

Attached is the *curriculum vitae* of Dr. Richard L. Boyd.

# **CURRICULUM VITAE**

# Richard Boyd

November 2007



| <b>Personal Details</b>        |  |              |         |
|--------------------------------|--|--------------|---------|
| Full name:                     | Richard Lennox Boyd  | Title: Profe | essor   |
| Current position:              | Director   |              |         |
| Department/School/<br>Section: | Monash Immunology and Stem Cell Laboratories (MISCL) / School of Biomedical Sciences |              |         |
| Faculty/Division:              | Medicine, Nursing and Health Sciences  | Campus:      | Clayton |
| Correspondence address:        | MISCL, Lvl 3, Bldg 75, Monash University, Wellington Rd, Clayton, 3800               |              |         |
| Email address:                 | richard.boyd@med.monash.edu.au   |              |         |

| Academic Qualifications Formal qualifications |                          |                   |  |  |
|---|--------------------------|-------------------|--|--|
| Year  | Qualification            | University        |  |  |
| 1972  | BSc (hons) – First class | Monash University |  |  |
| 1976  | PhD                      | Monash University |  |  |

## Appointments

| Year              | Position  | Location                    |
|-------------------|---|-----------------------------|
| Oct 2007- present | Director – Monash Immunology and Stem Cell Laboratories | Monash University - Clayton |
| 2005 – Sept 2007  | Deputy Director – MISCL                                 | Monash University - Clayton |
| 2003 - present    | Director – Immunology Program                           | Australian Stem Cell Centre |
| 2002 - present    | Chief Scientific Officer                                | Norwood Immunology Ltd      |

| 1995 - 2007           | Reader/Associate Professor – Professor | Department of Immunology, Monash University                         |  |
|-----------------------|--|---|--|
| Previous appointments |  |   |  |
| Year                  | Position                               | Location  |  |
| 1989-1994             | Senior Lecturer                        | Department of Pathology and Immunology,<br>Monash University.       |  |
| 1988                  | Lecturer (tenured appointment)         | Department of Pathology and Immunology, Monash University.          |  |
| 1985-1987             | Lecturer (fixed term)                  | Department of Pathology and Immunology,<br>Monash University.       |  |
| 1984                  | Research Fellow II                     | Department of Pathology and Immunology,<br>Monash University.       |  |
| 1983                  | Research Fellow I                      | Department of Pathology and Immunology,<br>Monash University.       |  |
| 1980-1982             | Research Fellow, Tenured position      | University of Innsbruck, Austria<br>(Universitats Vertragassistent) |  |
| 1978-1979             | Research Fellow                        | University of Innsbruck, Austria<br>(Universitats Vertragassistent) |  |
| 1976-1977             | Senior Tutor                           | Department of Pathology and Immunology,<br>Monash University.       |  |

| 2003 | Australian Federal Government Centenary Medal for Service to International Medical Research and Undergraduate Teaching  |
|------|---|
| 2004 | Australian Federal Government Business/Higher Education Round Table Award for Outstanding Achievement in Research and Development and Education and training.                   |
| 1981 | Preis der Stiftung der Hoechst Aktiengesellschaft, 1981  Effector mechanisms in spontaneous autoimmune thyroiditis in obese strain chickens. R.L. Boyd and G. Wick              |
| 1982 | Von Basedow-Forschungspreis, 1982  Effector mechanisms in the spontaneous autoimmune thyroiditis of obese strain chickens: analysis of cytotoxic cells.  R.L. Boyd and G. Wick. |

# Publications since 1995

| No. | Reference/Title of work   | Citation information |
|-----|---|----------------------|
| 1.  | Hince, M. N., Sakkal, S., Vlahos, K., Dudakov, J.A., Boyd, R.,* and Chidgey, A. C.* (2007) The role of sex steroids and gonadectomy in the control of   | IF: 1.709            |
| 2.  | thymic involution. <u>Cell. Immunol</u> (in press)  Chidgey, A.P, Dudakov, J.A., Seach, N., and <b>Boyd, R.L.</b> , (2007) Impact of niche aging on thymic regeneration and immune reconstitution. <u>Semin Immunol</u> , 19: 331-340 | IF: 10.00            |

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| ENCYCLOPEDIA OF LIFE SCIENCES. John Wiley & Sons, Ltd: Chichester (2007)  8. Rossi, S.W., Chidgey, A.P., Parnell, S.M., Jenkinson, W.E., Scott, H.S., Boyd, R.L., Lenkinson, E.J., and Anderson, G. (2007) Redefining epithelial progenitor potential in the developing thymus. Eur. J. Immunol. 37: 2411-2418.  6. Gray, D.H.D., Tull, D., Ueno, T., Seach, N., Classon, B.J., Chidgey, A.C., McConville, M., and Boyd, R.L., (2007) A Unique Thymic Fibroblast Population Revealed by the Monoclonal Antibody MTS-15. J. Immunol. 178: 4956-4965.  7. Goldberg, G., Alpdogan, O., Muriglan, S.J., Hammett, M., Milton, M.K., Eng., J. W., Hubbard, V.M., Kochman, A., Willis, L.M., Greenberg, A.S., Tjoe, K.H., Sutherland, J.S., Chidgey, A., van den Brink, M. and Boyd, R.L. (2007). Enhanced Immuno Reconstitution by Sex Steroid Abbation Following Allogeneic Hematopoietic Stem Cell Transplantation. J. Immunol. 178: 7473 – 7484.  8. Ann P. Chidgey and Richard L., Boyd (2006) Stemming the tide of thymic ageing. Nature Immunol. 7(10):1013-6.  9. Jeremy Millar, Jayne Sutherland, & Richard Boyd (2006) Alternative explanations for T-cell response to in-situ gene therapy for prostate cancer: In reply to Fugita et al (Int. J. Radiation Oncology Biol. Phys., vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., 66 (5): 1599.  10. Goldberg, G., Alpdogan, O., Muriglan, S.J., Hammett, M., Milton, M.K., Eng. J. M., Hubbard, V.M., Kochman, A., Willis, L.M., Greenberg, A.S., Tjoe, K.H., Sutherland, J.S., Chidgey, A., van den Brink, M. and Boyd, R.L. (2007). Enhanced Immune Reconstitution by Sex Seroid Abdiation Following Allogeneic Hematopoietic Stem Cell Transplantation. J. Immunol. (in press. M. No. 06-2215.).  11. Daniel H.D. Gray, Natalic Seach, Tomoo Ueno, Morag Milton, Adrian Liston, Andrew M. Lew, Christopher C. Goodnow, Richard L. Boyd (2006) Developmental kinetics, tunewer and stimulatory capacity of thymic epithelial cells. Blood 108(12):3777-85  12. Nijhof J.G. Braun K.M. Ginagreco A. van Pelt C. Kawamoto H. Boyd R.L. Willemar | 3.  | generation and regeneration: A new paradigm for establishing clinical   | IF: 9.422  |
| R.L., Jenkinson, E.J., and Anderson, G. (2007) Redefining epithelial progenitor potential in the developing thymos. Eur. J. Immunol. 37: 2411-2418.  6. Gray, D.H.D., Tull, D., Ueno, T., Seach, N., Classon, B.J., Chidgey, A.C., McConville, M., and Boyd, R.L., (2007) A Unique Thymic Fibroblast Population Revealed by the Monoclonal Antibody MTS-15. J. Immunol. 178: 4956-4965.  7. Goldberg, G., Alpdogan, O., Muriglan, S.J., Hammett, M., Milton, M.K., Eng., J. M., Hubbard, V.M., Kochman, A., Willis, L.M., Greenberg, A.S., Tjoe, K.H., Sutherland, J.S., Chidgey, A., van den Brink, M. and Boyd, R.L. (2007). Fahanced Immunol. 70: 1817-1819.  8. Ann P. Chidgey and Richard L. Boyd (2006) Stemming the tide of thymic ageing. Nature Immunol. 7(10):1013-6.  9. Jeremy Millar, Jayne Sutherland, & Richard Boyd (2006) Alternative explanations for T-cell response to in-situ gene therapy for prostate cancer: In reply to Fugita et al (In. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006. Int. J. Radiation On | 4.  | ENCYCLOPEDIA OF LIFE SCIENCES. John Wiley & Sons, Ltd:  | NA         |
| McConville, M., and Boyd, R.L., (2007) A Unique Thymic Fibroblast Population Revealed by the Monoclonal Antibody MTS-15. J. Immunol. 178: 4956-4965.  7. Goldberg, G., Alpdogan, O., Muriglan, S.J., Hammett, M., Milton, M.K., Eng, J. M., Hubbard, V.M., Kochman, A., Willis, L.M., Greenberg, A.S., Tjoc, K.H., Sutherland, J.S., Chidgey, A., van den Brink, M. and Boyd, R.L. (2007). Enhanced Immune Reconstitution by Sex Steroid Ablation Following Allogeneic Hematopoietic Stem Cell Transplantation. J. Immunol. 178: 7473 – 7484.  8. Ann P. Chidgey and Richard L. Boyd (2006) Stemming the tide of thymic ageing. Nature Immunol. 7(10):1013-6.  9. Jeremy Millar, Jayne Sutherland, & Richard Boyd (2006) Alternative explanations for T-cell response to in-situ gene therapy for prostate cancer: In reply to Fugine et al (Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 1, pp. 84-90, 2006). Int. J. Radiation Oncology Biol. Phys., 66 (5): 1599.  10. Goldberg, G., Alpdogan, O., Muriglan, S.J., Hammett, M., Milton, M.K., Eng, J. M., Hubbard, V.M., Kochman, A., Willis, L.M., Greenberg, A.S., Tjoc, K.H., Sutherland, J.S., Chidgey, A., van den Brink, M. and Boyd, R.L. (2007). Enhanced Immune Reconstitution by Sex Steroid Ablation Following Allogeneic Hematopoietic Stem Cell Transplantation. J. Immunol. (in press, Ms. No. 06-2215).)  11. Daniel H.D. Gray, Natalie Seach, Tomoo Ueno, Morag Milton, Adrian Liston, Andrew M. Lew, Christopher C. Goodnow, Richard L. Boyd (2006) Developmental kinetics, turnover and stimulatory capacity of thymic epithelial cells. Blood 108(12):3777-85  12. Nijhof JG. Braun KM. Giangreco A. van Pelt C. Kawamoto H. Boyd R.L. Willemze R. Mullenders LH. Watt FM. de Gruijl FR. van Ewijk W. (2006) Cheelopmental kinetics, turnover and stimulatory capacity of thymic epithelial cells. Blood 108(12):3777-85  13. Uldrich, A.P., Berzins, S.P., Malin, M.A., Bouillet, P., Strasser, A., Smyth, M.J., Boyd, R.L.* and Godfrey, D.I.* (2006). Antigen challenge inhibits  | 5.  | <b>R.L.</b> , Jenkinson, E.J., and Anderson, G. (2007) Redefining epithelial progenitor potential in the developing thymus. <u>Eur. J. Immunol</u> . 37:  | 1          |
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| 62. | Takeoka, Y., Chen, S-Y., <b>Boyd, R.L.</b> , Tsuneyama, K., Taguchi, N., Morita, S., Yago, H., Suehiro, S., Ansari, A., Schultz, L.D. and Gershwin, M.E.           | IF: NA     |
|     | (1997). A comparative analysis of the murine thymic microenvironment in normal, autoimmune and immunodeficiency states. <u>Dev. Immunol.</u> 5: 79 -               | CI: 11     |
| 62  | 89.  |            |
| 63. | Chidgey, A. and <b>Boyd</b> , <b>R.L.</b> (1997). Agonist peptide modulates T cell selection thresholds through qualitative and quantitative shifts in CD8 co-     | IF: 3.317  |
| 64. | receptor expression. <u>International Immunology</u> , 9: 1527 - 1536.   | CI: 31     |
| 04. | Chen, S-Y., Takeoka, Y., Pike-Nobile, L., Ansari, T., <b>Boyd, R.L.</b> and Gershwin, M.E. (1997). Autoantibody production and cytokine profiles of MHC class      | IF: NA     |
|     | I (β2-microglobulin) gene-deleted New Zealand Black (NZB) mice. <u>Clin.</u> <u>Immunol. Immunopathol.</u> 84: 318 327.  | CI: 7      |
| 65. | Jeurissen, S.H.M., Claassen, E. and <b>Boyd</b> , <b>R.L.</b> (1997). Immunohistochemistry in the analysis of lymphoid organs and their function. Methods in Avian | IF: NA     |
|     | Immunolgy 2235 - 2245.   |            |
| 66. | Siatskas, C., Lowenthal, J. and <b>Boyd, R.L</b> . (1997). Avian Cytokines. Methods in Avian Immunolgy. 2257 - 2268.   | IF: NA     |
|     | <u></u>  | CI: NA     |
| 67. | Hala, K., Malin, G., Dietrich, H., Loesch, U., Boeck, G., Wolf, H., Kaspers, B., Geryk, J., Falk, M., Wick, G. and Boyd, R.L. (1996). Analysis of the              | IF: 2.491  |
|     | initiation period of spontaneous autoimmune thyroiditis (SAT) in Obese strain (OS) chickens. <u>J. Autoimmunity</u> 9: 129 - 138.                                  | CI: 13     |
| 68. | Penit, C., Lucas, B., Vasseur, F., Rieker, T. and <b>Boyd, R.L.</b> (1996). Thymic   | IF: NA     |
|     | medulla epithelial cells acquire specific markers by post-mitotic maturation. <u>Dev.Immunol.</u> 5: 25 - 36.  | CI: 17     |
|     |  |            |

|            |   | ·          |
|------------|---|------------|
| 69.        | Chen, S-Y., Takeoka, Y., Ansari, T., <b>Boyd, R.L.</b> , Klinman, D. and Gershwin, E. (1996). The natural history of disease expression in CD4 and CD8 gene   | IF: 6.387  |
|            | deleted New Zealand Black mice. <u>J.Immunol.</u> 157: 2676 - 2684.   | CI: 17     |
| 70.        | Siatskas, C. McWaters, P.G., Digby, M. Lowenthal, J.W. and Boyd, R.L.   | IF: 3.261  |
|            | (1996). In vitro characterisation of a novel avian haemopoietic growth factor derived from stromal cells. <u>Dev.Comp.Immunol.</u> 20: 139 – 156.   | CI: 2      |
| 71.        | Blackburn, C.C., Augustine, C.L., Lui, R., Harvey, R.P., Malin M.A., Boyd,  | IF: 10.231 |
|            | <b>R.L.</b> , Miller, J.F.A.P. and Morahan, G. (1996). The nu gene acts cell-autonomously and is required for differentiation of thymic epithelial progenitors <u>Proc. Natl. Acad. Sci.</u> . <b>93</b> : 5742 - 5746.         | CI: 47     |
| 72.        | Boyd, R.L. and Siatskas, C. (1996). Improvement of chicken disease resistance by haemopoietic cytokines. Proc. Australian Poultry Science Symposium.  |            |
| 73.        | Takeoka, Y., Chen, S-Y., Yago, H., Boyd, R.L., Suehiro, S., Shultz, L.D.,   | IF: 2.201  |
|            | Ansari, T. and Gershwin, M.E. (1996). The murine thymic microenvironment changes with age. <u>Int. Archs. Allergy Appl. Immunol.</u> 111: 5 - 12.   | CI: 21     |
| 74.        | Obranovich, T.D. and <b>Boyd, R.L.</b> (1996) A bursal stromal derived cytokine   | IF: 3.261  |
|            | induces proliferation of MHC class II bearing cells. <u>Dev. Comp.</u> <u>Immunol.</u> 20: 61 - 75.   | CI: 4      |
|            |   |            |
| 75.        | Davey, G.D., Tucek-Szabo, C.L. and <b>Boyd</b> , <b>R.L.</b> (1996). Characterisation of the AKR thymic microenvironment and its influence on thymocyte   | IF: 2.372  |
|            | differentiation and lymphoma development. <u>Leukaemia Research</u> 20: 853  – 866  | CI: 2      |
| 76.        | Takeoka, Y., Whitmer, K.J., Chen, S-Y., Ansari, T., <b>Boyd, R.L.</b> , Schultz, L.D., Suehiro, S. and Gershwin, M.E. (1995). Thymic epithelial cell  | IF: NA     |
|            | abnormalities in (NZB x H-2 <sup>u</sup> ) F1 mice. <u>Clin. Immunol. Immunopathol.</u> <b>76</b> : 297 - 307.  | CI: 13     |
| 77.        | Duncan, M., Berman, B., Van de Water, J. Boyd, R.L., Wick, G. and   | IF: 2.201  |
|            | Gershwin, M.E.(1995).Mononuclear cells isolated from fibrotic skin lesions in avian scleroderma constitutively produce fibroblast-activating cytokines and immunoglobulin M. Int. Archs. Allergy Appl. Immunol. 107: 519 - 526. | CI: 3      |
| 78.        | Takeoka, Y., Yoshida, S.H., Morita, S., Suehiro, S., Boyd, R.L. and Gershwin,   | IF: NA     |
|            | M.E.(1995). Influence of neurotropin on thymic microenvironmental abnormalities of NZB mice. Int.J. Immunotherapy xi: 49 - 56.  | CI: 1      |
|            |   |            |
| 79.        | Takeoka, Y., Yoshida, S.H., van de Water, J., <b>Boyd, R.L.</b> , Suehiro, S., Ansari, T., and Gershwin, E. (1995) Thymic microenvironmental  | IF: 2.491  |
|            | abnormalities in MRL/MP-lpr/lpr, BXSB/MpJ Yaa and C3H/HeJ - gld / gld mice. <u>J. Autoimmunity</u> 8: 145 – 161   | CI: 14     |
| 80.        | Randle-Barrett, E.S. and Boyd, R.L. (1995). Thymic microenvironment and   | IF: NA     |
|            | lymphoid response to sublethal irradiation. <u>Dev. Immunol.</u> 4: 101 - 116.  | CI: 5      |
| BOOK CHAPT |   |            |
| 1.         | Gray, D.H.D., Gill, J.W., Trounson, A.O., and <b>Boyd, R.L.</b> (2004). Thymus and tolerance in transplantation. Handbook of Stem Cells, Volume 1, Elsevier, Academic Press. 675- 686   |            |
| 2.         | Hugo, P and Boyd, R. (2002). "Thymus". Encyclopeaedia of Life Sciences.p1-11  |            |
| 3.         | Boyd, R.L. (1998). Stromal cells. Encyclopaedia of Immunology 2233 – 2238   |            |
| L          |   |            |

| No   | Title of grant   | Funding body  | Year   | Successful | Individual contribution | Total value of<br>grant (\$)  |
|--|--|---|--|------------|-------------------------|---|
|  |  |   |  |            | %                       |   |
| Parameter Control of C | Australia – China Centre for Excellence in<br>Stem Cell Research. PI, Professor<br>Richard Boyd, Monash<br>Immunology and Stem Cell<br>Laboratories; Co-PI Professor<br>Lingsong Li Peking University<br>Stem Cell Research Centre | Australian Federal Government (DEST)  | 2007-<br>2010                                | YES        | 15                      | \$450,000<br>plus<br>\$450,000<br>cash/in kind<br>from Monash           |
| 2  | Innovative stem cell-based strategies to establish immune tolerance and tissue repair  | NH&MRC (Program grant)  | 2007-<br>2011                                | YES        | 30                      | \$5,338,070   |
| 3  | Reversal of multiple sclerosis-like disease by stem cell-based strategies  | National Multiple Sclerosis<br>Society, New York  | 2007-<br>2009                                | YES        | 20                      | Approx<br>\$360,000 US  |
| 4  | Novel generic vaccine approaches applied for the prevention of hepatitis C and influenza virus infections  | NH&MRC (Project grant)  | 2007-<br>2009                                | YES        | 20                      | \$376,875   |
| 5  | Therapeutic potential of fetal membrane derived stem cells in the treatment of chronic lung diseases   | Monash University Strategic<br>(Collaborative) grant  | 2007   | YES        | 15                      | \$60,000  |
| 6  | Rebuilding Immunity for Survival –<br>Immune Modulation Program  | Australian Stem Cell Centre   | 2006<br>2007                                 | YES        | 80                      | ~\$1mil<br>~\$1mil  |
| 7  | Research and development and pre-clinical and clinical trials on the effectiveness of LHRH agonists on rejuvenating T cell-based immune responses.   | Norwood Immunology  | 2006<br>2005<br>2004<br>2003<br>2002<br>2001 | YES        | 80                      | ~\$500k<br>~\$1.7 mil<br>~\$1.7 mil<br>~\$1.3 mil<br>~\$904k<br>~\$478k |
| 8  | Rebuilding Health for survival   | National Institutes for Health (USA) CA100265-01  Boyd research-based Phase II randomised placebo controlled FDA approved clinical trial. | 2004<br>2005<br>2006<br>2007                 | YES        | 10                      | No direct funds to lab  |
| 9  | Cellular molecular and functional analysis of thymic epithelial precursor cells  | National Institutes for Health (USA) RO1. AI 57477  | 2004<br>2005<br>2006<br>2007                 | YES        | 10                      | \$250,000 p.a.<br>No direct<br>funds to lab                             |
| 10   | Strategies To Prevent and Repair Thymic Microenvironment   | National Institutes for Health, (USA) RO1   | 2004<br>2005<br>2006<br>2007                 | YES        | 10                      | \$250,000 p.a.<br>No direct<br>funds to lab                             |

| 11 | The use of Neural Stem Cells as a  Therapeutic Tool in Neurological Disorders              | Baker Foundation   | 2006<br>2007<br>2008 | YES | 30  | \$300k<br>\$300k<br>\$300k                 |
|----|--|--|----------------------|-----|-----|--|
| 12 | Molecular mechanism of the thymus organogenesis workshops                                  | Australian Government Department of Education, Science and Training. (International Science Linkages Strategic Policy)(SP0500002)                      | 2005<br>2006         | YES | 100 | \$45k total                                |
| 13 | Pathogenesis of clostridial myonecrosis  | NH&MRC (Project)<br>(now NH&MRC Program Grant<br>ID 284214 for 5 years, J.Rood<br>principal investigator. Boyd<br>associated research ~\$120,000 p.a.) | 2003                 | YES | 10  | \$130k                                     |
| 14 | The generation, fate and functional  | NH&MRC (Project)   | 2000                 | YES | 50  | \$105,231                                  |
|    | potential of recent thymic emigrants   |  | 2001                 |     |     | \$103,705                                  |
|    |  |  | 2002                 |     |     | \$103,705                                  |
| 15 | The cellular and molecular basis to the  | NH&MRC (Project)   | 2000                 | YES | 80  | \$88,327                                   |
|    | paradox of positive versus negative T cell selection.                                      |  | 2001                 |     |     | \$93,142                                   |
|    |  |  | 2002                 |     |     | \$94,223                                   |
| 16 | The molecular basis to the development   | NH&MRC (Project)   | 2001                 | YES | 80  | \$75,000                                   |
|    | and function of the thymic microenvironment.   |  | 2002                 |     |     | \$75,000                                   |
|    |  |  | 2003                 |     |     | \$75,000                                   |
| 17 | Differential gene expression analysis to elucidate the mechanisms of                       | NH&MRC (Project)   | 1998                 | YES | 50  | \$47,325                                   |
|    | thymic development.  |  | 1999                 |     |     | \$50,083                                   |
|    |  |  | 2000                 |     |     | \$52,882                                   |
| 18 | Evaluation of Diver Health   | Victorian Fisheries  | 1997                 | YES | 50  | \$15,000                                   |
|    |  |  | 1998                 |     |     |  |
| 19 | Molecular Studies of Lymphostromal Molecules Important in Thymopoiesis and T Cell Function | NH&MRC (Project)   | 1997<br>1998<br>1999 | YES | 80  | \$90,495.49<br>\$93,343.60<br>\$94,333.89  |
| 20 | Cellular and Molecular Basis to Intrathymic Positive Selection                             | NH&MRC (Project)   | 1997<br>1998<br>1999 | YES | 80  | \$62,104.22<br>\$63,714.17<br>\$65, 321.81 |
| 21 | Functional Analysis of the Thymic Microenvironment   | NH&MRC (Project)   | 1997<br>1998<br>1999 | YES |     | \$58,040.22<br>\$59,650.17<br>\$61, 257.81 |
| 22 | Generation of Antigen-restricted Human T cells in vitro                                    | Alfred HealthCare Group  | 1996<br>1997         | YES | 20  | \$20,000 pa                                |
| 23 | The Thymic Microenvironment in NZB mice.   | National Institutes for Health,<br>Bethesda  | 1996<br>1997<br>1998 | YES | 20  | ~\$110,000 p.a.                            |

| 24 | Development of an in vitro assay for positive selection (Site visit to Ontario Cancer Institute, AMGEN, Toronto) | Department of Industry, Technology and Regional Development. | 1996                         | YES | 50  | \$7,000  |
|----|--|--|------------------------------|-----|-----|--|
| 25 | Improvement of Chicken Disease<br>Resistance by Cytokines  | Australian Chicken Meat<br>Research Council                  | 1995<br>1996<br>1997         | YES | 100 | \$30,000<br>\$30,000<br>\$30,000                 |
| 26 | Molecular analysis of novel lymphostromal antigens regulating thymopoiesis.                                      | NH&MRC (Project)   | 1995<br>1996<br>1997         | YES | 80  | \$29,540<br>\$30,012<br>\$30,012                 |
| 27 | Immunoregulatory properties of whey proteins   | Dairy Research Development Council.                          | 1995<br>1996<br>1997         | YES | 100 | \$29, 500<br>\$29, 500<br>\$29, 500              |
| 28 | Immunoregulation in Avian Scleroderma  | National Institutes of Health,<br>USA                        | 1993<br>1994<br>1995<br>1996 | YES | 20  | \$121,629<br>\$121,629<br>\$121,629<br>\$121,629 |

| Assess | Assessor of Competitive Grants for:                               |  |  |  |  |  |
|--------|---|--|--|--|--|--|
| No.    | Funding body (i.e. ARC / NHMRC / Other)                           |  |  |  |  |  |
| 1      | NH&MRC, Australia   |  |  |  |  |  |
| 2      | Australian Research Council (ARC)                                 |  |  |  |  |  |
| 3      | Australian Stem Cell Center                                       |  |  |  |  |  |
| 4      | Anti-Cancer Councils of Victoria, New South Wales, and Queensland |  |  |  |  |  |
| 5      | Human Frontier Science Program                                    |  |  |  |  |  |
| 6      | MRC United Kingdom  |  |  |  |  |  |
| 7      | National Institutes of Health, Bethesda                           |  |  |  |  |  |
| 8      | Alfred Hospital, Melbourne  |  |  |  |  |  |
| 9      | New Zealand HRC   |  |  |  |  |  |
| 10     | Raine Foundation, Western Australia                               |  |  |  |  |  |
| 11     | USA-Israel Binational Science Foundation                          |  |  |  |  |  |
| 12     | Diabetes Vaccine Development Centre                               |  |  |  |  |  |
| TOTAL  | Approximately 5- grant reviews per year                           |  |  |  |  |  |

| Compl | Supervisory Ro   |           |
|-------|--|-----------|
| 1.    | Gabrielle Goldberg (2006) "Sex steroid ablation enhances immune reconstitution following severe immunodepletion"                             | MAIN      |
| 2.    | Tamara Etto (2006) Generation of minor histocompatibility antigen specific T cells using various antigen sources for adoptive immunotherapy. | ASSOCIATE |
| 3.    | Kate Ward (2006) Key steps involved in the generation of antigen specific T cells for adaptive immunotherapy.                                | ASSOCIATE |
| 4.    | Tracy Heng (2005) "Kinetics and functional impact of castration-induced thymic regeneration."  | MAIN      |
| 5.    | Adam Uldrich (2004) "The effect of antigen encounter on T cell export, recent thymic emigrants, and NKT cells."                              | MAIN      |
| 6.    | David Zammit (2003) "Development and Immunological characterisation of mice deficient in Thymic Shared Antigen-1."                           | MAIN      |

| 7.  | Bryce Feltis (2003) "Eosinophils and eosinophilic chemokines in asthma and the effect of inhaled corticosteroids."                                  | ASSOCIATE |
|-----|---|-----------|
| 8.  | Daniel Gray (2003) "Thymic stromal cells: Population dynamics an their role in thymopoiesis."   | JOINT/CO  |
| 9.  | Jason Gill (2002) "Progenitor cells of the thymic epithelium, and factors influencing thymic development."  | MAIN      |
| 10. | Jayne Sutherland (2001) "Enhancement of thymopoiesis and cell-mediated immunity through sex steroid blockade."                                      | MAIN      |
| 11. | Stuart Berzins (2000) "The development of thymic emigrants and their influence on the peripheral T cell pool."                                      | MAIN      |
| 12. | Christopher Siatskas (1998) "Identification and functional characterisation of chicken cytokines."  | MAIN      |
| 13. | Ann Chidgey (1998) "The cellular and molecular basis to thymic positive selection."   | MAIN      |
| 14. | Oskar Lechner (1997) "Endogenous glucocorticoids in autoimmune disease and in the thymus."  | MAIN      |
| 15. | Marina Katerelos (1997) "Isolation and characterisation of a cyclosporin A-downregulated gene: CsA-19."   | MAIN      |
| 16. | John Emmins (1996) "Immunological and genetic analysis of Koalas."  | MAIN      |
| 17. | Gayle Davey (1995) Characterisation of a lymphostromal molecule involved in normal and leukaemic thymopoiesis."                                     | MAIN      |
| 18. | Marilina Masciantonio (1995). "Characterisation of anti-thymic stromal mAbs and the antigens they detect."  | MAIN      |
| 19. | Elise Randle (1994). "Functional evaluation of the lymphostromal molecule TSA-1."   | MAIN      |
| 20. | Andrew Bean (1993). "Chicken Lymphocyte and Stromal Cell-associated Antigens: A phenotypic and Functional Analysis."                                | MAIN      |
| 21. | Tania Obranovich (1993). "The Bursal Microenvironment: Production and Phenotypic and Functional Characterisation of Stromal Cell Lines."            | MAIN      |
| 22. | Natalie J. Davidson (1992). "Phenotypic and Functional Analysis of the Chicken Thymic Microenvironment."  | MAIN      |
| 23. | Carolyn L. Tucek (1991) "Analysis of Antigens Present on Both Mouse Thymic Stromal Cells and Thymocytes: Relevance to Thymopoiesis".                | MAIN      |
| 24. | Trevor J. Wilson (1990) "Chicken Thymic and Bursal Stromal Cells: Characterisation, Ontogeny and Alterations in Immunodeficiency and Autoimmunity." | MAIN      |
| 25. | Dale I. Godfrey (1990) "Thymic Stromal Microenvironments: Characterisation, Modulation and Functional Analysis."                                    | MAIN      |
| 26. | Gary A.Waanders, (1990) "Modulation of T Cell Differentiation by Cytokine and Monoclonal Antibody Treatment of Fetal Thymic Organ Culture."         | MAIN      |
| 27. | Anthony Eglezos (1989) "Neurogenic Modulation of the Immune Response."  | MAIN      |
| 28. | Jeanette A.Tratkiewicz (1988) "Cytotoxic Cells in Acute Leukaemia: Diagnostic and Therapeutic Potential."   | JOINT/CO  |
| 29. | Helen C. Ramm (1989) "Chicken B Lymphocyte Development: Ontogeny, Bursal Microenvironment and Alterations in Infectious Bursal Disease."            | MAIN      |
| 30. | Geoffrey W. Dandie (1988) "Antigenic Promotion, Antigenic Competition and Immunity." Cosupervised with Dr D. Poskitt.                               | JOINT/CO  |

| 31. David J. Izon (1989) "The Role of Thymic Stromal Cells in Lymphopoiesis and Myelopoiesis".                  | MAIN      |
|---|-----------|
| 32. Ian G. Barr (1984) "Leucocyte Differentiation and Surface Markers in Man and Mouse."                        | JOINT/CO  |
| Completed MSc   |           |
| 33. Maree Hammett (2002) "T cells and the Graft-versus leukaemia phenomenon."                                   | JOINT/CO  |
| 34. Elizabeth O'Flaherty (1994) "Assessment of bone marrow stromal cell function from patients with leukaemia." | JOINT/CO  |
| Current Research Students - PhD   |           |
| 35. Linda Kenins  | MAIN      |
| 36. Anne Fletcher   | MAIN      |
| 37. Jarrod Dudakov  | MAIN      |
| 38. Samy Sakal  | MAIN      |
| 39. Jessica Reiseger  | MAIN      |
| 40. Adele Barnard   | MAIN      |
| 41. Katerina Vlahos   | MAIN      |
| 42. Natalie Seach   | MAIN      |
| 43. Melanie Hince   | MAIN      |
| 44. Joanna Lim  | JOINT     |
| 45. Sarah Snelgrove   | JOINT     |
| 46. Chew –Li Soh  | JOINT     |
| 47. Oanh Ngyen  | ASSOCIATE |

| Year   | Course       | Student            |
|--------|--------------|--------------------|
| 2006/7 | BMSc (Hons)  | Danika Khong       |
| 2006/7 | BSc (Hons)   | Harry Banaharis    |
| 2006   | BMSc (Hons)  | Mirabelle Ho       |
| 2006   | BMSc (Hons)  | Jordan Kane        |
| 2006   | BSc (Hons)   | Tamara Lowen       |
| 2005   | BSc (Hons)   | Melanie Hince      |
| 2005   | BMSci (Hons) | Timothy Williams   |
| 2004   | BSc (Hons)   | Marc Jurblum       |
| 2003   | BSc (Hons)   | Jessica Porter     |
| 2003   | BSc (Hons)   | Natalie Seach      |
| 2003   | BSc (Hons)   | Katerina Vlahos    |
| 2003   | BSc (Hons)   | Jarrod Dudakov     |
| 2002   | BSc (Hons)   | Samantha Harris    |
| 2002   | BSc (Hons)   | Lauren Young       |
| 2002   | BSc (Hons)   | Anne Fletcher      |
| 2001   | BSc (Hons)   | Elizabeth Chapman  |
| 2001   | BSc (Hons)   | Alexander Barker   |
| 2001   | BSc (Hons)   | Samy Sakkal        |
| 2000   | BSc (Hons)   | Gerard Tarulli     |
| 2000   | BSc (Hons)   | Tracy Heng         |
| 1999   | BSc (Hons)   | Adam Uldrich       |
| 1999   | BSc (Hons)   | Douglas Liddicoat  |
| 1999   | BSc (Hons)   | Gabrielle Goldberg |
| 1998   | BSc (Hons)   | Daniel Pellici     |
| 1998   | BSc (Hons)   | Morag Milton       |
| 1998   | BSc (Hons)   | Daniel Gray        |
| 1997   | BSc (Hons)   | David Zammit       |
| 1997   | BSc (Hons)   | Maree Hammett      |
| 1997   | GDip (Imm)   | Bryce Feltis       |
| 1997   | BMSci(Hons)  | Oskar Lechner      |
| 1997   | BSc (Hons)   | Rebecca Baird      |
| 1997   | BSc (Hons)   |                    |
|        |              | Jayne Sutherland   |
| 1996   | BSc (Hons)   | Jason Gill         |
| 1996   | BSc (Hons)   | Darren Ellemor     |
| 1995   | BMSci (Hons) | Sanjay Swaminathan |
| 1995   | Gdip (Imm)   | Hieng Lim          |

| 1994 | BSc (Hons)   | Julian Pearson        |
|------|--------------|-----------------------|
| 1993 | BSc (Hons)   | Melinda Price         |
| 1993 | BSc (Hons)   | Ann Chidgey           |
| 1992 | BSc (Hons)   | Stuart Fraser         |
| 1991 | BSc (Hons)   | Mirey Lahoud          |
| 1990 | BSc (Hons)   | Christopher Siatskas  |
| 1990 | BSc (Hons)   | Elise Randle          |
| 1989 | BSc (Hons)   | Uta Bauer             |
| 1989 | BSc (Hons)   | Belinda Gourlay       |
| 1989 | MSc (Prelim) | Mark Malin            |
| 1987 | BSc (Hons)   | Frank Sommerfeld      |
| 1987 | BSc (Hons)   | Carolyn Tucek         |
| 1987 | BSc (Hons)   | Sara Gipton           |
| 1987 | BSc (Hons)   | Marilina Masciantonio |
| 1987 | BSc (Hons)   | Natalie Davidson      |
| 1986 | BSc (Hons)   | Dale Godfrey          |
| 1986 | BSc (Hons)   | Anthony Eglezos       |
| 1986 | BSc (Hons)   | Andrew Bean           |
| 1985 | BMSci (Hons) | Georg Oberhuber       |
| 1985 | BSc (Hons)   | Gary Waanders         |
| 1985 | BSc (Hons)   | Kathy Mitrangas       |
| 1985 | BSc (Hons)   | Lisa Bonadonna        |
| 1985 | BSc (Hons)   | Paul Bello            |
| 1984 | BSc (Hons)   | Nicholas Samaras      |
| 1984 | BSc (Hons)   | Dianne Young          |
| 1984 | BSc (Hons)   | Helen Ramm            |
| 1983 | BSc (Hons)   | Trevor Wilson         |
| 1983 | BSc (Hons)   | David Izon            |
| 1983 | BSc (Hons)   | Brendan Classon       |
| 1977 | BSc (Hons)   | Gregory Woods         |
| 1977 | BSc (Hons)   | lan G Barr            |

| Unit (Subject)<br>code | Unit Name                       | Sessions<br>(Lecture, tutorial,<br>practical etc.) | No. of students    | Years<br>taught  | Improvement/innovation   |
|------------------------|---------------------------------|--|--------------------|------------------|--|
| IMM3011                | Immunology                      | Lectures and<br>Practical Classes                  | 50-100<br>per year | 1976-<br>present | Co-founded Immunology as a Science subject. Involved in the continual evolution of the the subject – lecture content, practical classes, evaluations.  |
| IMM4000                | Immunology –<br>honours subject | Research Projects including theory component       | 15-25 per<br>year  | 1976-<br>present | Co-founder of subject. Involved in the continual evolution of the subject, including acting as coordinator for 8 of 16 years since 1990. Organised extensive overhaul of course examination. |

| Editorial and Scientific Appraisal Services |  |                |  |  |  |
|---|--|----------------|--|--|--|
| Editorial duties                            | Publication                              | Years          |  |  |  |
| Editor-in-chief                             | Developmental Immunology                 | 1999 - 2003    |  |  |  |
| Member – editorial board                    | Developmental and Comparative Immunology | 1993 - 2003    |  |  |  |
| Member – editorial board                    | Clinical and Developmental Immunology    | 2004 - present |  |  |  |
| Reviewer                                    | Autoimmunity                             | Ongoing        |  |  |  |
| Reviewer                                    | Blood                                    | Ongoing        |  |  |  |
| Reviewer                                    | Experimental Biology and Medicine        | Ongoing        |  |  |  |
| Reviewer                                    | European Journal of Immunology           | Ongoing        |  |  |  |
| Reviewer                                    | Immunity                                 | Ongoing        |  |  |  |
| Reviewer                                    | Immunology and Cell Biology              | Ongoing        |  |  |  |
| Reviewer                                    | Journal of Autoimmunity                  | Ongoing        |  |  |  |
| Reviewer                                    | Journal of Clinical Investigation        | Ongoing        |  |  |  |
| Reviewer                                    | Journal of Experimental Medicine         | Ongoing        |  |  |  |

| TOTAL:   | Approx: 35-40 manuscript reviews per year |         |
|--|---|---------|
| Reviewer   | Journal of Experimental Gerontology       | Ongoing |
| Reviewer   | Trends in Immunology                      | Ongoing |
| Reviewer Proceedings of the National Academy of Sciences |   | Ongoing |
| Reviewer Nature Reviews Immunology                       |   | Ongoing |
| Reviewer   | Nature Immunology                         | Ongoing |
| Reviewer   | ver Nature                                |         |
| Reviewer   | eviewer Journal of Immunology             |         |

| Role   | Committee   | Years                                  |  |
|--|---|--|--|
| Co-convenor and member,<br>Scientific advisory board | Discovery, Science and Biotechnology conference, Melbourne.   | 2006                                   |  |
| Chairman   | Organising Committee – "Rebuilding Immunity for Better Health – an interface between immunology, stem cells, and regenerative medicine."  | 2005                                   |  |
| Member   | Scientific committee – 15th International conference on lymphatic tissues and germinal centres in immune reactions. Potsdarm. Germany.  | 2005                                   |  |
| Member   | Scientific Organising Committee, 2nd Australian Stem<br>Cell Conference, Sydney.  | 2004                                   |  |
| Member   | Scientific Committee, 14th International conference on lymphatic tissues and germinal centres in immune reactions. Groningen.   | 2002                                   |  |
| Co-chairman  | Organising Committee, Australian Society for Immunology Annual General Meeting, Melbourne.  | 1998                                   |  |
| Co-chairman  | Organising Committee for International Conference on<br>Germinal Centres and Lymphatic Tissues in<br>Immune Reactions, Graz, Austria.   | 1996                                   |  |
| Chairman   | Organising and Scientific Committee, International Workshop on T Lymphocytes, Heron Island. ThymOz I Oct 11 -15, 1995, ThymOz II March 25 - 29, 1998, ThymOz III April 11-18, 2000, ThymOz IV April 1-5, 2003, ThymOv V April 5-10, 2006. | 1995, 1998, 2000,<br>2003, 2006.       |  |
| Member   | Scientific and Organising Committees, Foundation member, International Workshop on the Thymus. Rolduc, The Netherlands.1988, 1991, 1995, 1997, 2001, 2004.  | 1988, 1991, 1995,<br>1997, 2001, 2004. |  |
| Member   | Avian Leucocyte CD nomenclature committee   | 1990-2000                              |  |
| Member   | Organising Committee for Australian Society for Immunology Annual Scientific Meeting, Melbourne.  | 1990                                   |  |
| Member   | Cluster of Thymic Epithelial Staining (CTES) nomenclature committee   | 1989-0995                              |  |

| Member | Organising Committee, German Society for Immunology, Innsbruck. | 1979 |
|--------|---|------|
|        |   |      |

#### Invitations to conferences (since 1995)

**2007** Invited speaker, Pennington Scientific Symposium, "NeuroImmune Signaling and Inflammation," Baton Rouge, Louisiana, USA

Invited speaker, Discovery, Science and Biotechnology (DSB), Brisbane, Australia

Invited speaker, 1st Australia - China Biomedical Research Conference, Melbourne, Australia.

Invited plenary speaker, 5th ISSCR Annual Meeting, Cairns, QLD, Australia.

Invited speaker, Rolduc Workshop on Thymocyte and T cell Biology, The Netherlands

Invited speaker, Translating Basic HIV Immunology into Novel Interventions Workshop, Garvan Institute of Medical Research, Sydney. Part of 4th IAS Conference on HIV Pathogenesis, Treatment and Prevention.

Invited speaker, Australian Clinical Immunologists and Allergists (ASCIA) Annual Scientific Meeting, Freemantle, WA.

Invited speaker, 3rd Barossa Meeting "Signalling Systems," Barossa Valley, SA.

2006 Invited presenter "Immune based Therapies Working group" of the National Centre in HIV Epidemiology and Clinical Research (University of New South Wales).

Co-convenor and invited speaker – Discovery, Science and Biotechnology, Melbourne.

Invited speaker, Collaborative Planning Meeting between Monash University and the University of Newcastle upon Tyne, Prato.

Invited Presentation, Australian Stem Cell Centre's International Scientific Advisory Board meeting, Toronto.

Invited Lecturer, Science Week Public Lecture Series, Federation Square, Melbourne.

Invited speaker, 2nd Annual meeting of the Monash Infection and Immunity Network, Marysville

Invited Chairperson, Aging Research in Immunology (ARIG) symposium, Paris.

Invited speaker, Australian Stem Cell Centre workshop, "Ethics and Stem Cells – "Stem Cell Research Post the Lockhart Review." AH&MRC Congress, Melbourne.

Invited participant, Monash University delegation to Shanghai Institute for Biological Sciences (SIBS).

Invited speaker, Becton Dickinson National Sales Conference, Fiji.

#### 2005 Invited Speaker, Lorne Cancer Conference, Phillip Island, Victoria, February

Invited Speaker, 8th International Symposium on GnRH Analogues in Cancer and Human Reproduction, February, Salzburg, Austria.

Invited Speaker, Kyoto T Cell Conference, April 6-10, Kyoto, Japan

Invited Speaker, Japanese Society for the Promotion of Science, Core-to-Core Workshop on Thymus Organogenesis' April 11-12. Tokushima, Japan.

Invited Speaker, 3rd Congress of Federation of Immunology Societies of Asia-Oceania (FIMSA), Hangzhou, April 18-22.

Invited Speaker and Scientific Committee member, 15th International conference on lymphatic tissues and Germinal Centres in Immune Reactions . April 20-24 Potsdam, Germany

Invited Speaker Advances in Agricultural and Medical Biotechnology, 29-30 September Kuala Lumpur.

Invited Speaker, 6th Australian Peptide Conference, October 9-14, Hamilton Island, Queensland.

Invited Speaker, 10th Annual Australian Autoimmunity, 19-21st October.

Invited Speaker, Science in the Vines, November, Barossa Valley. South Australia

Invited Symposium Speaker. 35th Annual Meeting Australasian Society for Immunology. December 4-8 Melbourne.

Invited Lecturer, Post-graduate Teaching Workshop, 35th Annual Meeting Australasian Society for Immunology. December 4-8 Melbourne.

2004 Invited Symposium speaker. Swiss Society for Allergy and Immunology, Geneva April 15-17

Invited Speaker, Rolduc International T cell Workshop, Rolduc. The Netherlands. May 1-5.

Invited speaker, Japanese Society for the Promotion of Science, Core-to-Core Workshop on Thymus Organogenesis'. Tokushima August 16-18.

Invited Speaker, ThymUs - International Workshop on the Thymus. Peurto Rico, November.

Invited Speaker and Symposium Chairperson, 2nd Australian Stem Cell Conference/ Australian Health and medical Research Congress, November, Sydney.

Invited Symposium Speaker, British Society for Immunology, December 10th, Harrogate. UK.

Invited Symposium Speaker, Australian Society for Immunology, Adelaide, December

**2003** Invited speaker and corporate presentation Bio 2003, Washington June 22-27.

Invited Speaker, Workshop chairman, 1st National Stem cell Conference, Melbourne Oct 9-11.

Invited Speaker, First Barossa Meeting (Science amongst the Vines), "Regeneration". Nov 19-21.

Invited Symposium Speaker, Japanese Society for Immunology, December, 7-10.

2002 Invited symposium speaker, Kyoto T Cell Conf. (KTCC), Kyoto, Japan April 3-5.

Invited Symposium Speaker and Chairman, 14th International Conference on Lymphoid Tissues and Germinal Centres in Immune Reactions, Groningen, The Netherlands, June 23-27.

Invited Symposium Speaker, Federation of Immunological Societies of Australasia, Beijing China, October.

Invited key Note Speaker, Australian Society of Medicine, Canberra, Nov 21-23.

2001 Invited Symposium speaker. Thymus Workshop. Rolduc. Kerkrade. The Netherlands, May 12-15.2001.

Invited Speaker: IgV Meeting, Mt Buffalo, Victoria, October 14-16.

Invited Keynote Address, ThymUS International thymus meeting, Peurto Rico, November 2-6.

Invited Chairperson, ThymUS International Thymus Meeting, Peurto Rico, November 2-6.

Invited Chairperson, Australian Society for Immunology, Canberra. December 1-4.

**2000** Invited Symposium speaker and Chairperson, VII International Society Avian Endocrinology, Varanasi, India, Feb 1-5.

Invited Symposium speaker, XXI World Poultry Congress, Montreal, August 20-24.

Invited Chairperson. Australian Society for Immunology, Sydney. Dec.

**1999** Invited Speaker, FASEB/Clinical Immunology. Washington April 19-21.

Invited Speaker, Rolduc International Workshop on Thymus, May 1-4.

Invited Speaker, Austrian Academy of Science Special Symposium in Honour of G. Wick, Innsbruck, May 8.

Invited Symposium Speaker, 6th Asia/Oceania Regional Congress of Gerontology, June 8-11.

Invited Speaker and Workshop Chairman, 13th International Conference on Lymphoid Tissues and Germinal Centres in Immune Reactions, Geneva, Switzerland 1-6.

1998 Plenary Chairperson, symposium speaker, 25th Annual Meeting Australian Soc Immunology, Melbouren Nov 29-Dec3.

1997 Invited Chairperson 5th Conference of the Immunology Group of Victoria, Mt. Buffalo, March 16-18.

Invited speaker, International Workshop on Thymus, Rolduc/Kerkrade Holland, March 23-26.

Invited Symposium speaker 5th Kyoto T Cell Meeting, Kyoto, October 2-4.

Invited Symposium speaker and Chairperson, XIIIth Int Congress of Comparative. Endocrinology, Yokoma Nov 17-21.

**1996** Invited Symposium Speaker, Australian Poultry Science Symposium, Sydney, Feb 6-7.

Invited Symposium Speaker, International Symposium on Avian Endocrinology, Alberta, Canada, March 31 - April 5.

Invited Keynote Speaker, Avian Immunology Workshop, Obergurgl, Austria, April 21 –24.

Invited Symposium Speaker, 5th International Expert Forum on Immunotherapy and Gene Therapy, Jerusalem, June 4-9, 1996.

Invited Symposium Speaker and Workshop Chairman, 12th International Conference on Lymphoid Tissues and Germinal Centres in Immune Reactions, Graz, Austria, July 1-5, 1996.

Invited Plenary Session Speaker, First Congress of the Federation of Immunological Societies of Asia-Oceania, (ASI) Adelaide, Dec 1-5.

1995 Invited Co-chairman, 9th International Congress of Immunology, San Fransisco, California. USA, July.

#### **TOTAL INVITATIONS PRIOR TO 1995: 16**

## Patents (as at November 2007)

| Application No          | Title   | Inventor   | Status                                     |
|-------------------------|---|--|--|
| Provisional Application | Novel thymic cellular populations and uses thereof  | Anne Fletcher, Ann Chidgey,<br>Natalie Seach, Richard Boyd   |  |
| PCT/AU00/00329          | Improvement of T cell mediated immunity Filed on October 13, 2000 as United States Application Serial No. 09/795,302. Previously filed as: PP9778, Australia, April 15, 1999. PCT/AU/00329, Australia, April 17, 2000, PRO745, Australia, October 13, 2000. | Richard Lennox Boyd  | Granted 4 countries Pending 6 countries    |
| PCT/AU01/01291          | Treatment of T cell disorders   | Richard Lennox Boyd  | Granted 4 countries<br>Pending 9 countries |
| PCT/IB01/02739          | Haematopoietic stem cell gene therapy   | Richard Lennox Boyd  | Granted 2 countries<br>Pending 7 countries |
| PCT/IB01/02740          | Improvement of graft acceptance through manipulation of thymic regeneration   | Richard Lennox Boyd  | Granted 2 countries Pending 7 countries    |
| PCT/IB01/02352          | Normalisation of defective T cell responsiveness through manipulation of thymic regeneration  | Richard Lennox Boyd  | Granted 1 country Pending 6 countries      |
| PCT/IB01/02350          | Stimulation of thymus for vaccine development   | Richard Lennox Boyd  | Granted 3 countries Pending 6 countries    |
| PCT/IB01/02351          | Diagnostic indicator of thymic function   | Richard Lennox Boyd<br>Ann Patricia Chidgey  | Granted 1 country Pending 7 countries      |
| PCT/IB01/02745          | Disease prevention by reactivation if the thymus  | Richard Lennox Boyd  | Granted 3 countries Pending 8 countries    |
| PCT/AU03/00749          | Thymic epithelial cells with progenitor capacity  | Richard Lennox Boyd  | Lapsed                                     |
| PCT/US2004/011919       | Tolerance to graft prior to thymic regeneration   | Richard Lennox Boyd Gabrielle Lianne Goldberg Jayne Suzanne Sutherland Ann Patricia Chidgey                        | Pending 10 countries                       |
| PCT/US2004/011913       | Disease prevention and vaccination following thymic reactivation  | Norwood Immunology Ltd Richard Lennox Boyd Gabrielle Lianne Goldberg Jayne Suzanne Sutherland Ann Patricia Chidgey | Pending 1 country                          |
| PCT/US2004/011921       | Disease prevention and vaccination prior to thymic reactivation   | Norwood Immunology Ltd Richard Lennox Boyd Gabrielle Lianne Goldberg Jayne Suzanne Sutherland Ann Patricia Chidgey | Granted 1 country Pending 9 countries      |
| PCT/US2004/011920       | Tolerance to graft following thymic reactivation  | Richard Lennox Boyd Gabrielle Lianne Goldberg Jayne Suzanne Sutherland Ann Patricia Chidgey                        | Pending 1 country                          |

#### APPENDIX 1:

#### **PUBLICATIONS PRIOR TO 1995**

- 1. **Boyd, R.L.**, Rolland, J.M. and Cauchi, M.N. (1974) Membrane antigenic changes associated with PHA transformation of mouse spleen cells *in vitro*. Immunolog. Commun., **3**: 337-349.
- 2. **Boyd, R.L.**, Ward, H.A. and Muller, H.K. (1976) Antisera specific for the reticulin of the bursa of Fabricius. <u>Int. Archs of Allergy and Appl. Immunol.</u>, **50**:129-132.
- 3. Clarke, G.N., **Boyd, R.L.** and Muller, H.K. (1977) Actin-like protein in human sperm heads. In <u>"Immunological Influence of Human Fertility"</u> Ed. B. Boettcher, (Academic Press, Sydney, New York, London), pp. 211-214.
- 4. **Boyd, R.L.**, Toh, B.H., Muller, H.K. and Ward, H.A. (1977) Actin-like protein in chicken and mammalian lymphoid tissue demonstrated by reactivity with human smooth muscle autoantibody. <u>Int. Arch. Allergy Appl. Immunol.</u> 55:283-292.
- 5. **Boyd, R.L.** and Ward, H.A. (1978) Lymphoid antigenic determinants of the chicken: cellular representation and tissue localization. Immunology. **34**:9-17.
- 6. **Boyd, R.L.** and Ward, H.A. (1978) Antigenic changes associated with chicken B lymphocyte development. In "Advances in Experimental Medicine and Biology", 114 Eds. W. Muller-Ruchholtz and H.K. Muller-Hermelink. (Plenum Press, New York), pp.25-30.
- 7. Wick, G., Kofler, R., Gundolf, R., Muller, P.U. and **Boyd, R.L.** (1979) The nature of effector cells in experimental and spontaneous autoimmune thyroiditis. <u>In 6<sup>th</sup> International Convocation on Immunology</u>: "<u>Immunopathology</u>", Eds. F. Milgrom and B. Albini, (S. Karger Verlag Basel) pp. 101-106.
- 8. Wick, G. and **Boyd**, R.L. (1979) Effector mechanisms in thyroid autoimmune diseases. In: <u>Autoimmunity in Thyroid Diseases</u>, Eds. E. Klein and F.A. Forster, (F.K.Schattauer Verlag, Stuttgart, New York) pp. 23-29.
- 9. **Boyd**, R.L. and Ward, H.A. (1979) Induction of B- and T-cell differentiation by chicken bursa and thymus reticular epithelial cells. In "Cell Biology and Immunology of Leucocyte Function". pp. 235-239.
- 10. **Boyd**, R.L., Barr, I.G., Ward, H.A. and Muller, H.K. (1979) Antigeneic and functional properties of bursal and thymic reticular epithelial cells. Folia Biol. **25**:310-312.
- 11. **Boyd, R**. and Wick, G. (1979) Effector mechanisms in spontaneous autoimmune thyroiditis of Obese strain chickens. <u>Folia Biol.</u> **25**:340-342.
- 12. **Boyd**, R. and Wick G. (1979) Role of suppressor cells in autoimmune diseases. In <u>European Surgical Research</u>. Ed. W. Brendl, (S. Karger Verlag, Basel) pp.212-213.
- 13. Boyd, R. and Ward H.A. (1979) Antigenic changes associated with chicken B lymphocyte differentiation. Immunobiol. 156:186.
- 14. **Boyd**, R., Schauenstein, K. and Wick, G. (1979) Characterization of effector cells in spontaneous autoimmune thyroiditis. Immunobiol. **156**:244.
- 15. **Boyd, R.L.** and Wick, G. (1980) Killer cells in the chicken: a microcytotoxicity assay using antigen-coated erythrocytes as targets. <u>J.Immunol. Methods.</u> **35**: 233-247.
- 16. Wick, G. and **Boyd**, R. (1980) Effector and suppressor cells in Obese strain (OS) chickens with spontaneous autoimmune thyroiditis. Fed. Proc. **39**: 570.
- 17. **Boyd, R.** and Wick, G (1980) Effector mechanisms in spontaneous autoimmune thyroiditis in Obese strain chickens. In: <a href="Proceedings of the Serono Symposia">Proceedings of the Serono Symposia</a>; Ed. D. Doniach, G.F. Fenzi and L. Bachieri. Academic Press, London, New York, pp.199-206.
- 18. **Boyd, R.L.** and Wick, G. (1981) Ontogenic development of cytotoxic and suppressor cells in Obese strain (OS) chickens with spontaneous thyroiditis. In: <u>Aspects of Developmental and Comparative Immunoloy I.</u> Ed. J.B. Solomon (Pergamon Press, Oxford, New York) pp. 543-544.

- 19. **Boyd, R.L.**, Barr, I.G., Ward, H.A. and Muller H.K. (1980) Functional and antigenic analysis of the bursal and thymic microenvironments. In: <u>Aspects of Developmental and Comparative Immunology 1.</u> Ed. J.B. Solomon, (Pergamon Press, Oxford, New York) pp.537-538.
- 20. Wick, G., **Boyd, R.L.**, Hala, K., de Carvalho, L., Muller, P.U. and Cole, R.K. (1981) The Obese strain (OS) of chickens with spontaneous autoimmune thyroiditis. Review of recent data. <u>Curr. Top. Microbiol. and Immunol.</u> **91**: 110-128.
- 21. Wick, G., **Boyd, R.L.**, de Carvalho, L. and Roitt, I.M. (1981). Analysis of suppressor cells in chickens of the Obese strain (OS) with spontaneous autoimmune thyroiditis. In: <u>Cellular and Molecular Mechanisms of Immunologic Tolerance.</u> Ed. T. Hraba and M. Hasek. Marcel Dekker, Inc. New York, Basel pp. 501-506.
- 22. Wick, G., **Boyd**, **R.L.** and de Carvalho, L. (1981) The role of T cells in spontaneous autoimmune thyroiditis in OS chickens. <u>Fed.</u> Proc. **40**: 1140.
- 23. Hala, K., **Boyd**, **R.L.** and Wick, G. (1982) Chicken major histo-compatibility complex and disease (review). <u>Scand J. Immunol.</u> 14: 607-616.
- 24. Wick, G., **Boyd, R.L.**, Hala, K., Thunold, S. and Kofler, H. (1982) Pathogenesis of spontaneous autoimmune thyroiditis in Obese strain (OS) chickens. <u>Clin. Exp. Immunol.</u> 47: 1-18.
- 25. Thunold, S., **Boyd**, **R**., Schauenstein, K. and Wick, G. (1982) Tissue localization of lymphocyte surface antigens and receptors for IgG-Fc and C' in the chicken. J. Histochem. Cytochem. **30**: 201-206.
- 26. Woods, G.M. and **Boyd, R.L.** (1982) Contact-dependent expression of actin in chicken lymphocytes <u>in vitro</u>. <u>Int. Arch. Allergy Appl. Immunol.</u> **67**: 335-339.
- 27. **Boyd, R.L.** and Wick, G. (1982) Effector mechanisms in the spontaneous autoimmune thyroiditis of Obese strain (OS) chickens: Analysis of cytotoxic cells. Int. Arch. Allergy. Immunol. **69**: 286-295.
- 28. Wick, G., Schauenstein, K., Thunold, S. and **Boyd, R.L.** (1982) IgG-Fc and C3 receptors in the chickens: distribution tissue localization and functional significance. Molec. Immunol. 19: 1267-1273.
- 29. Barr, I.G., Alderton, M.R., Brumley, J.L., **Boyd, R.L.**, Muller, H.K. and Ward H.A. (1982) Antigens associated with bursal and thymic reticular epithelial cells. <u>Advances in Experimental Medicine</u> and Biology. **149**: 711-717.
- 30. Wick, G., **Boyd**, **R.L.** and Muller, P.U. (1982) The effect of Cyclosporin A on spontaneous autoimmune thyroiditis in the Obese strain (OS) of chickens. <u>Advances in Experimental Medicine</u> and Biology. **149**: 19-23.
- 31. **Boyd, R.L.**, Hala, K., Boyd, J. and Wick, G. (1982) Quantitative analysis and interactions of immunoregulatory cells in the chicken thymus. <u>Immunobiol.</u> 163: 156.
- 32. Traill, K.N., Bock, G., **Boyd, R.L.** and Wick, G. (1983) Chicken thrombocytes. Isolation, serological and functional characterization using the fluorescence activated cell sorter. Dev. Comp. Immunol. 7: 111-125.
- 33. **Boyd**, **R.L.**, Ward, H.A. and Muller, H.K. (1983) Bursal and thymic reticular epithelial cells in the chicken: Preparation of <u>in vitro</u> monolayer cultures. <u>J. Retic. Soc.</u> **34**: 371-382.
- 34. **Boyd, R.L.**, Ward, H.A. and Muller, H.K. (1983) Bursal and thymic reticular epithelial cells in the chicken: Induction of B and T lymphocyte differentiation by <u>in vitro</u> monolayer cultures. J. Retic. Soc. 34: 383-393.
- 35. **Boyd, R.L.** and Wick, G. (1983) Genetically controlled degree of autoimmune thyroiditis in Obese strain (OS) chickens is expressed at both the humoral and cellular level. <u>Immunol. Commun.</u> 12: 263-272.
- 36. **Boyd, R.L.** and Wick, G. (1983) Autoimmune thyroiditis of Obese strain (OS) chickens. <u>Immunopathologie</u>: <u>Collection Foundation</u> Marcel Merieux pp. 113-119.
- 37. **Boyd**, **R.L.** and Ward, H.A. (1984) Lymphoid antigenic determinants of the chicken: ontogeny of bursa-dependent lymphoid tissue. <u>Dev. Comp. Immunol.</u> **8**: 149-167.
- 38. **Boyd, R.L.**, Oberhuber, G., Hala, K. and Wick, G. (1984) Obese strain (OS) chickens with spontaneous autoimmune thyroiditis have a deficiency in thymic nurse cells. J. Immunology, **132**: 718-724.

- 39. Hala, K., **Boyd, R.L.**, Wolf, H., Bock, G. and Wick, G. (1984) Functional analysis of B-L (Ia-like) antigen bearing chicken peripheral blood cells. <u>Scand. J. Immunol.</u> **20**: 15-19.
- 40. Wolf, H., Hala, K., **Boyd, R.L.** and Wick, G. (1984) MHC and non-MHC-encoded surface antigens of chicken lymphoid cells and erythrocytes recognized by polyclonal xeno-, allo- and monoclonal antibodies. <u>Eur. J. Immunol.</u> 14: 831-839.
- 41. Wick, G., Hala, K., Wolf, H., **Boyd, R.L.** and Schauenstein, K. (1984) Distribution and functional analysis of B-L/Ia positive cells in the chicken: expression of B-L/Ia antigens on thyroid epithelial cells in spontaneous autoimmune thyroiditis. <u>Molec. Immunol.</u> **12**: 1259-1265.
- 42. Hala, K., Wick, G., **Boyd, R.L.**, Wolf, H., Bock, G. and Ewert, D.L. (1984) The B-L (Ia-like) antigens of the chicken, lymphocyte plasma membrane distribution and tissue localization. Dev. Comp. Immunol. **8**: 673-682.
- 43. Traill, K.N., Bock, G., **Boyd**, **R.L.**, Ratheiser, K. and Wick, G. (1984). Ontogeny of surface markers on functionally distinct T cell subsets in the chicken. <u>Eur. J. Immunol.</u> 14: 61-67.
- 44. Wick, G., Oberhuber, G., **Boyd, R.L.** and Hala, K. (1984). Obese strain (OS) chickens with spontaneous autoimmune thyroiditis have a deficiency in thymic nurse cells. In: <u>Lymphoid cell function in Ageing.</u> Ed. A.L. de Weck. <u>Topics in Aging Research in Europe, Vol. 3. EURAGE Rijswijk 1984, pp. 39-46.</u>
- 45. Andrews, P. and **Boyd, R.L.** (1985). The murine thymic nurse cell: an isolated thymic microenvironment. <u>Eur. J. Immunol.</u> **15**: 36-42.
- 46. **Boyd, R.L.**, Hala, K. and Wick, G. (1985). Interactions and quantitative analysis of immunoregulatory cells in the chicken thymus. J. Immunol. 135: 3039-3049.
- 47. Andrews, P., **Boyd**, **R.L.** and Shortman, K. (1985). The limited immunocompetence within murine thymic nurse cells. <u>Eur. J.</u> <u>Immunol</u>. **15**: 1043-1048.
- 48. Scollay, R., Andrews, P., **Boyd**, **R**. and Shortman, K. (1985) The role of the thymic cortex and medulla in T cell differentiation. <u>Adv. Exp. Med. Biol.</u> **186**: 229-234.
- 49. Shortman, K., Scollay, R., Wilson, A., Andrews, P., **Boyd, R.**, Butcher, E., Weissman, I. (1986). Mature and immature thymocytes: surface phenotype, immune function and intrathymic location. <u>Prog. in Leukocyte Biology Vol. 5.</u> (Eds. Oppenheim, J.J. and Jacobs, D.M.), pp. 3-10.
- 50. Shortman, K., Scollay, R., Andrews, P. and **Boyd, R.L.** (1986). Development of T lymphocytes within the thymus and within thymic nurse cells. <u>Current Topics Microbiol. Immunol.</u> **126**: 5-18.
- 51. **Boyd, R.L.**, Wilson, T.J., Mitrangas, K. and Ward, H.A. (1987) Characterization of chicken thymic and bursal stromal cells. In: Avian Immunology II, Eds W.T.Weber and D.L. Ewert; Alan R. Liss Inc. New York, N.Y. pp 29-39.
- 52. **Boyd, R.L.**, Mitrangas, K., Ramm, H.C., Wilson, T.J., Fahey, K.J. and Ward, H.A. (1987) Chicken B lymphocyte differentiation: ontogeny, bursal microenvironment and effect of IBD virus. In: <u>Avian Immunology</u> II. Eds. W.T. Weber and D.L. Ewert; Alan R. Liss Inc. New York, N.Y. pp. 41-51.
- 53. Helme, R.D., Eglezos, A., Dandie, G.W., Andrews, P.V. and **Boyd, R.L.** (1987). The effect of Substance P on the regional lymph node antibody response to antigenic stimulation in capsaicin pre-treated rats. <u>J. Immunol.</u> **139**:3470-3473.
- 54. Eglezos, A., Helme, R.D., Dandie, G.W., Andrews, P.V. and Boyd, R.L. (1988). Substance P-Mediated modulation of the primary antibody response. Adv. Exp. Med. Biol. 237: 499-503.
- 55. Godfrey, D.I., Izon, D.J., Wilson, T.J., Tucek, C.L. and **Boyd, R.L.** (1989). Ontogeny, <u>in vitro</u> culture and modulation by immunosuppression <u>in vivo</u> of thymic stromal elements defined by M.Abs. Adv. Exp. Med. Biol. **237**: 269-276
- 56. Tratkiewicz, J.A., Szer, J. and Boyd, R.L. (1989). Are LAK cells a possible form of immunotherapy? Adv. Exp. Med. Biol. 237: 447-450
- 57. Tratkiewicz, J.A., Szer, J. and **Boyd**, **R.L.** (1989). Blood Leukocyte NK activity as an early indicator of leukaemic relapse. <u>Adv. Exp. Med. Biol.</u> **237**: 451-456
- 58. Ramm, H.C., Mitrangas, K.H., Wilson, T.J., Boyd, R.L. and Ward, H.A. (1989). Chicken B lymphocyte differentiation ontogeny

- 59. Wilson, T.J., Mitrangas, K.H., Ramm, H.C., **Boyd, R.L.** and Ward, H.A. (1989). Response of chicken bursa stroma to treatment with cyclophosphamide and IBD virus. <u>Adv. Exp. Med. Biol.</u> **237**: 75-8
- 60. Shortman, K., Vremec, D., D'Amico, A., Battye, F. and **Boyd, R.L.** (1989). Nature of the thymocytes associated with dendritic cells and macrophages in thymic rosettes. <u>Cell Immunol</u>. **119**: 85-100
- 61. Tucek, C.L., **Boyd, R.L.** and Hiai, H. (1989) Antigens shared by thymic stromal cells and T lymphocytes are abnormally expressed in AKR thymuses. Thymus 14: 95-107.
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#### **ATTACHMENT B**

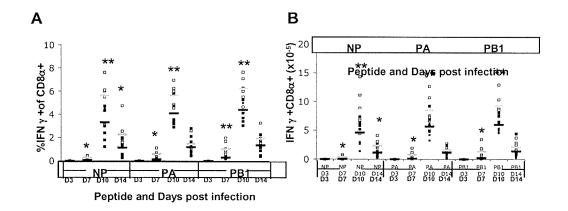


FIGURE.1 Decreases in the response time and magnitude of the primary response to the NP<sub>366</sub>, PA<sub>224</sub> and PB<sub>1703</sub> epitopes in the spleen of aged mice (black squares, mean = black bar) compared to young mice (open square, mean = grey bar). Naïve mice were infected with Hkx31 influenza A virus, and groups of 5 to 6 were sampled 3,7 10 and 14 days later. CD8 $\alpha$ + enriched splenocytes were stimulated for 5 h with 1mM NP366 , PA224 or PB1703 peptide and analysed for expression of CD8 $\alpha$ , IFN $\gamma$ , TNF $\alpha$  and IL2. The percentage (A) and numbers of virus-specific CD8 $\alpha$ + IFN $\gamma$  expressing cells (B) are shown. Numbers were calculated using cell counts and the percentage of cells staining. The data presented depict the responses of 5 to 6 mice from two or three experiments. Statistical significance was determined using an unpaired two-tailed Student's t test (\*, p<0.05; \*\*,p<0.01).

#### **ATTACHMENT C**

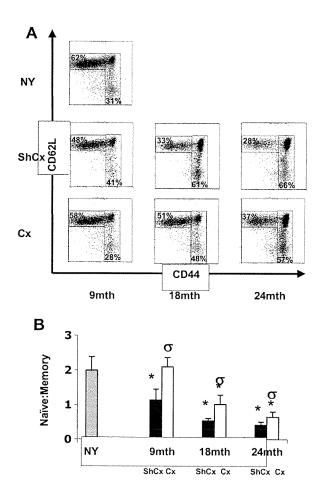


FIGURE 2. Decreases in the naïve to memory ratio with age and the reversal seen with sex steroid ablation. Enriched splenocytes were stained with CD62L and CD44 (A) to obtain a ratio of naïve to memory T cells (B) from NY (grey), 9mth, 18mth and 24mth old that had either been ShCx (black) or Cx (white) 6 weeks prior. Representative facs plots with percentage of naïve (CD62L+CD44-) and memory (CD62L±CD44+) CD8α+ cells shown in bold (A). Statistical significance was determined using an unpaired two-tailed Student's t test (for NY versus ShamCx and Cx •, p<0.01; for ShamCx vs Cx σ, p<0.05).

#### **ATTACHMENT D**

Note: IFNy+CD8+ cells and NP+ cells are virus specific cells

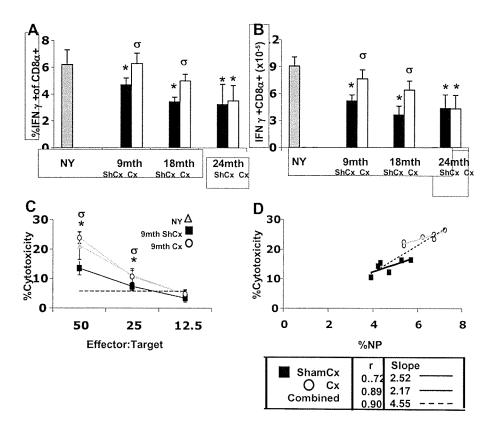


FIGURE 3. Decreases in the magnitude of the primary response to the NP<sub>366</sub> in aged ShCx mice (black) compared to normal young mice (NY) (grey) and the reversal of this in aged Cx mice (white). Normal young and 9mth, 18mth and 24mth (ShCx or Cx) naïve mice were infected with Hkx31 influenza A virus, and groups of 5 to 6 were sampled 10 days later. CD8α and IFNy expression was analysed as described in Fig.1. CD8α and IFNγ expression was analysed as described in Fig1. Percentage (A) and number (B) of CD8 $\alpha$ + IFN $\gamma$  expressing cells are shown for the spleen. The cytotoxicity of splenocytes from normal young (grey triangles) and 9mth old ShCx (black squares) or Cx (open circles) mice were tested against NP<sub>366</sub>-pulsed EL4 target cells. The values shown are the means and ±SD from 5 to 6 individual mice (C). Correlations between cytotoxic activity (E:T of 50:1) and DbNP<sub>366</sub> tetramer staining is shown for 9mth old (ShCx and Cx) individual mice (D). The data presented depict the responses of 5 to 6 mice from two or three experiments. Statistical significance was determined using an unpaired two-tailed Student's t test (for NY versus ShamCx or Cx •, p<0.05; for ShamCx vs Cx  $\sigma$ , p<0.05).

#### **ATTACHMENT E**

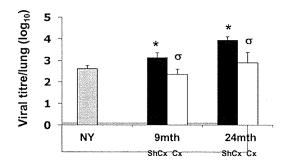


FIGURE 4. Increased viral titres in aged ShCx mice (black) compared to normal young mice (NY (grey)) and the reversal of this following Cx (white). Viral titres were measured (as described in Fig. 3) from NY, 9mth ShCx/Cx and 24mth ShCx/Cx mice 7 days post primary infection. The data presented depict the responses of 5 to 6 mice from two or three experiments. Statistical significance was determined using an unpaired two-tailed Student's t test (for NY versus ShamCx •, p<0.05; for ShamCx vs Cx σ, p<0.05).